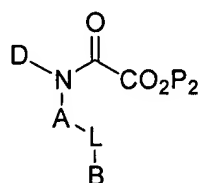


Amendments to the Claims

1 (Currently Amended).

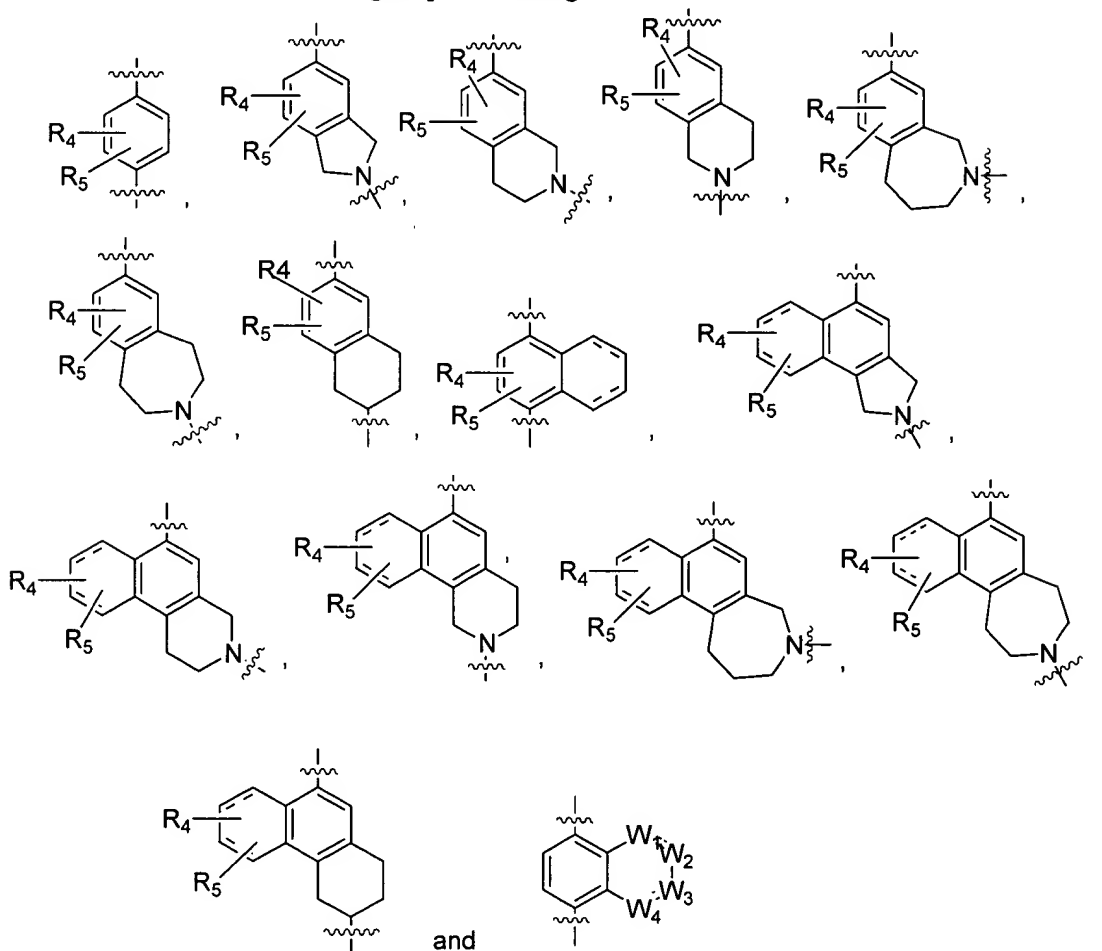
A compound of formula (I)



(I),

or a therapeutically acceptable salt or prodrug thereof, wherein

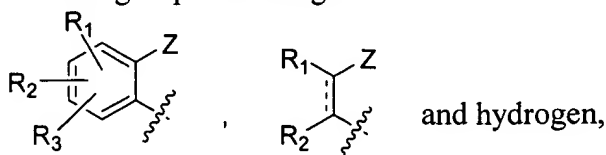
A is selected from the group consisting of



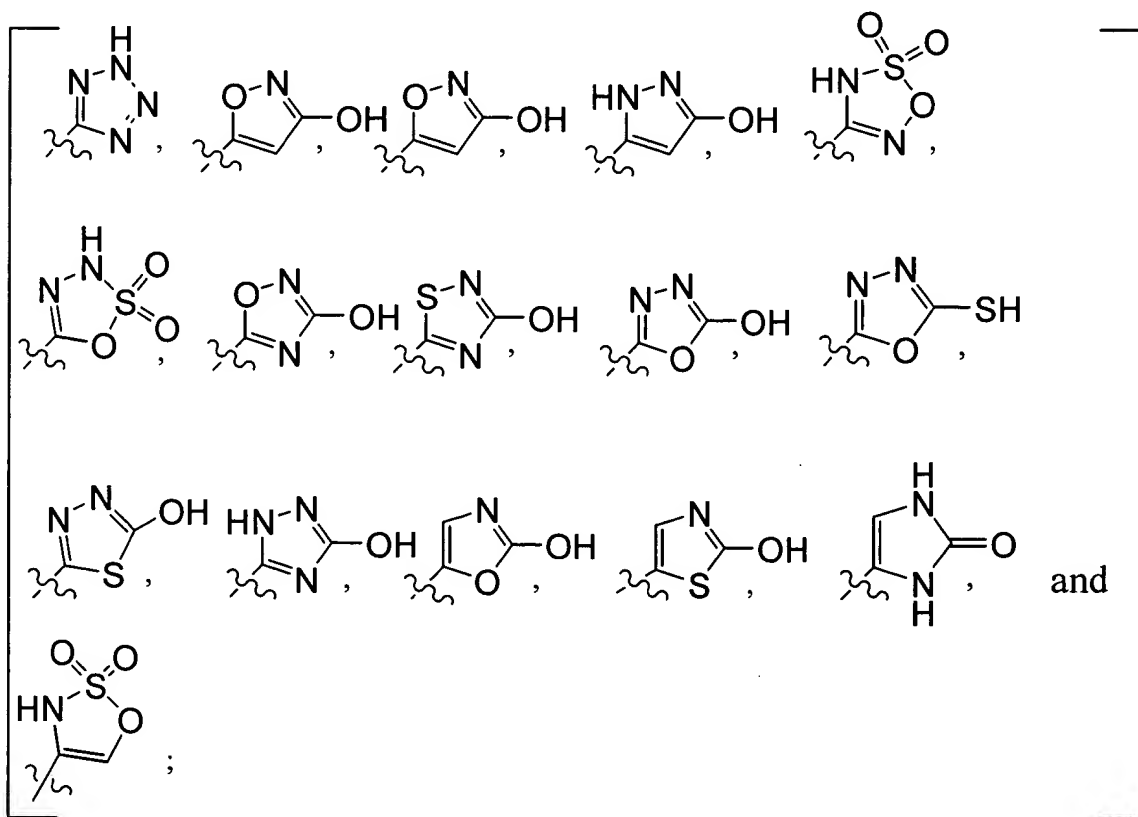
wherein the dotted line is either absent or is a single bond;

B is selected from the group consisting of hydrogen, alkyl, aryl, arylalkyl, ~~heterocycle and heterocyclealkyl;~~

D is selected from the group consisting of



wherein Z is selected from the group consisting of alkoxy, alkyl, amino, cyano, nitro, CO_2P_1 , SO_3H , $\text{PO}(\text{OH})_2$, $\text{CH}_2\text{PO}(\text{OH})_2$, $\text{CHFPO}(\text{OH})_2$, $\text{CF}_2(\text{PO}(\text{OH})_2)$, and $\text{C}(\text{=NH})\text{NH}_2$; **and the following 5-membered heterocycles:**



wherein P_1 and P_2 are independently selected from the group consisting of hydrogen, alkyl, alkenyl, arylalkyl, cycloalkyl and (cycloalkyl)alkyl;

R_1 , R_2 , R_3 , R_4 and R_5 are independently selected from the group consisting of hydrogen, alkoxy, alkyl, aryl, arylalkyl, cyano, halo, haloalkoxy, haloalkyl, **heterocycle**, **heterocyclealkyl**, hydroxy, hydroxyalkyl, nitro, $\text{NR}_\text{A}\text{R}_\text{B}$, $\text{NR}_\text{A}\text{R}_\text{B}\text{C}(\text{O})$, $\text{NR}_\text{A}\text{R}_\text{B}\text{C}(\text{O})\text{alkyl}$ and $\text{NR}_\text{A}\text{R}_\text{B}\text{C}(\text{O})\text{alkenyl}$, wherein R_A and R_B are independently selected from the group consisting of hydrogen, alkyl, alkoxy, carbonyl, alkylsulfonyl, aryl, arylalkyl, carbonyl, arylcarbonyl, arylsulfonyl and $(\text{R}_\text{C}\text{R}_\text{D})\text{carbonyl}$ wherein R_C and R_D are independently selected from the group consisting of hydrogen, alkyl, aryl, and arylalkyl, **or R_A and R_B taken together with the nitrogen to which they are attached form a ring selected from the group consisting of pyrrolidine, piperidine, morpholine, homopiperidine and piperazine;**

L is selected from the group consisting of

- $-(\text{CH}_2)_m\text{X}_1(\text{CH}_2)_n\text{CH}(\text{R}_8)\text{C}(\text{R}_9\text{A})(\text{R}_9\text{B})\text{X}_2(\text{CH}_2)_p\text{C}(\text{O})\text{N}(\text{R}_{10})\text{CH}(\text{CO}_2\text{R}_{11})(\text{CH}_2)_q\text{X}_3^-$;
- $-(\text{CH}_2)_m\text{X}_1(\text{CH}_2)_n\text{CH}(\text{R}_8)\text{C}(\text{R}_9\text{A})(\text{R}_9\text{B})\text{X}_2(\text{CH}_2)_p\text{EC}(\text{O})\text{N}(\text{R}_{10})\text{CH}(\text{CO}_2\text{R}_{11})(\text{CH}_2)_q\text{X}_3^-$;
- $-(\text{CH}_2)_m\text{X}_1(\text{CH}_2)_n\text{CH}(\text{R}_8)\text{C}(\text{R}_9\text{A})(\text{R}_9\text{B})\text{X}_2(\text{CH}_2)_p\text{X}_3^-$;
- $-(\text{CH}_2)_m\text{X}_1(\text{CH}_2)_n\text{CH}(\text{R}_8)\text{C}(\text{R}_9\text{A})(\text{R}_9\text{B})\text{X}_2(\text{CH}_2)_p\text{X}_3(\text{CH}_2)_q\text{X}_4^-$; and

$-(CH_2)_mX_1(CH_2)_nCH(R_8)C(R_{9A})(R_{9B})X_2(CH_2)_pE(CH_2)_qX_3-$, wherein each group is drawn with the left end attached to A and the right end attached to B;

m, n, p and q are independently between 0-4;

R_8 is selected from the group consisting of hydrogen, hydroxy, NR_AR_B and $(NR_AR_B)alkyl$;

R_{9A} and R_{9B} are independently selected from the group consisting of hydrogen, alkyl, hydroxyalkyl and $R_ER_FNalkyl$, wherein R_E and R_F are independently selected from the group consisting of hydrogen, alkyl, alkoxycarbonyl and alkanoyl, or R_{9A} and R_{9B} taken together are oxo;

R_{10} is selected from the group consisting of hydrogen, alkyl, alkanoyl and alkoxycarbonyl;

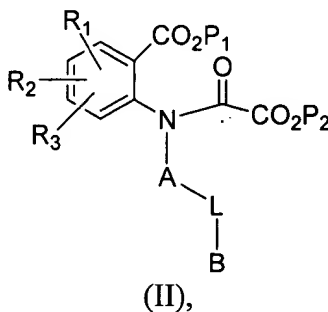
R_{11} is independently selected from the group consisting of hydrogen, alkyl, alkenyl, arylalkyl, cycloalkyl, and (cycloalkyl)alkyl;

E is selected from the group consisting of aryl and cycloalkyl;

X_1 , X_2 , X_3 , and X_4 are independently absent or are independently selected from the group consisting of NR_G , O, S, $S(O)$ and $S(O)_2$, wherein R_G is selected from the group consisting of hydrogen, alkyl, alkanoyl and alkoxycarbonyl; and

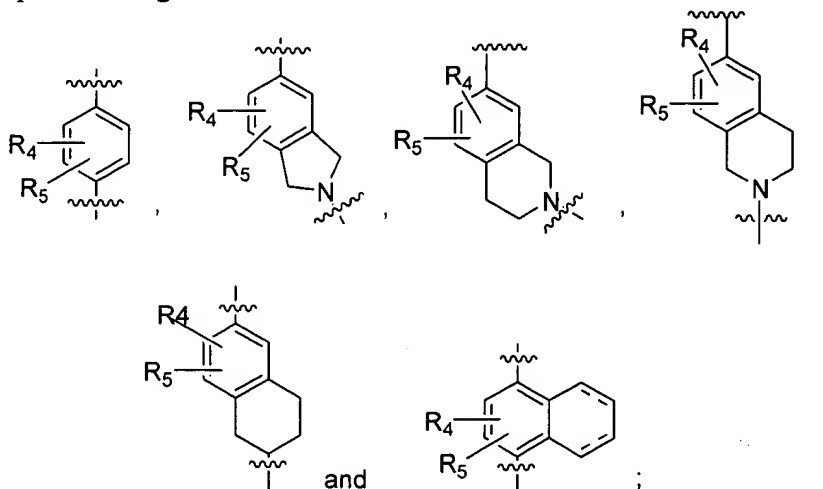
W_1 , W_2 , W_3 and W_4 are independently selected from the group consisting of CH, CH_2 , N, NH and O.

2 (Original). The compound according to claim 1 of formula (II)



or a therapeutically acceptable salt or prodrug thereof wherein A, B, L, P_1 , P_2 , R_1 , R_2 , and R_3 are defined in Claim 1.

3 (Currently Amended). The compound according to claim 2, wherein A is selected from the group consisting of



R_1 , R_2 , R_3 , R_4 and R_5 are independently selected from the group consisting of hydrogen, alkoxy, alkyl, cyano, halo, haloalkoxy, haloalkyl, ~~heterocycle~~, hydroxy, hydroxyalkyl, nitro, NR_AR_B , $NR_AR_BC(O)$, $NR_AR_BC(O)alkyl$ and $NR_AR_BC(O)alkenyl$;

R_{10} is selected from the group consisting of hydrogen and alkyl; and

R_{11} is independently selected from the group consisting of hydrogen, alkyl and arylalkyl.

4 (Original). The compound according to claim 2, wherein

L is

$-(CH_2)_mX_1(CH_2)_nCH(R_8)C(R_{9A})(R_{9B})X_2(CH_2)_pC(O)N(R_{10})CH(CO_2R_{11})(CH_2)_qX_3-$.

5 (Original). The compound according to claim 2, wherein

L is

$-(CH_2)_mX_1(CH_2)_nCH(R_8)C(R_{9A})(R_{9B})X_2(CH_2)_pC(O)N(R_{10})CH(CO_2R_{11})(CH_2)_qX_3-$; and R_8 is NR_AR_B .

6 (Original). The compound according to claim 2, wherein

L is

$-(CH_2)_mX_1(CH_2)_nCH(R_8)C(R_{9A})(R_{9B})X_2(CH_2)_pC(O)N(R_{10})CH(CO_2R_{11})(CH_2)_qX_3-$; R_8 is NR_AR_B ; and R_{9A} and R_{9B} together are oxo.

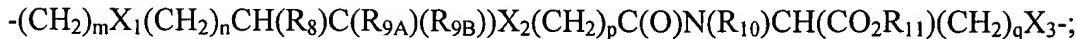
7 (Original). The compound according to claim 2, wherein

L is

$-(CH_2)_mX_1(CH_2)_nCH(R_8)C(R_{9A})(R_{9B}))X_2(CH_2)_pC(O)N(R_{10})CH(CO_2R_{11})(CH_2)_qX_3-$; R_8 is NR_AR_B ; R_{9A} and R_{9B} together are oxo; and X_2 is NR_C .

8 (Currently Amended). The compound according to claim 2, wherein

L is



R_8 is $\text{NR}_\text{A}\text{R}_\text{B}$;

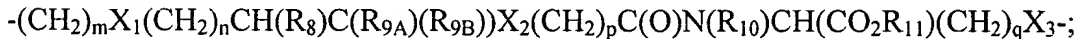
$\text{R}_{9\text{A}}$ and $\text{R}_{9\text{B}}$ together are oxo;

X_2 is NR_C ; and

B is ~~selected from the group consisting of~~ aryl ~~and heterocycle~~.

9 (Currently Amended). The compound according to claim 2, wherein

L is



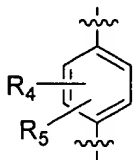
R_8 is $\text{NR}_\text{A}\text{R}_\text{B}$;

$\text{R}_{9\text{A}}$ and $\text{R}_{9\text{B}}$ together are oxo;

X_2 is NR_C ;

B is ~~selected from the group consisting of~~ aryl ~~and heterocycle~~; and

A is

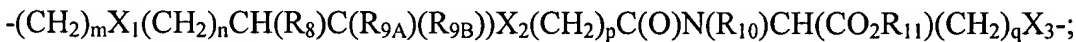


10 (Original). The compound according to claim 9, which is

N-[5-({*N*-acetyl-4-[(carboxycarbonyl)(2-carboxyphenyl)amino]-3-ethylphenylalanyl}amino)pentanoyl]-*L*-tyrosine.

11 (Original). The compound according to claim 2, wherein

L is



R_8 is $\text{NR}_\text{A}\text{R}_\text{B}$;

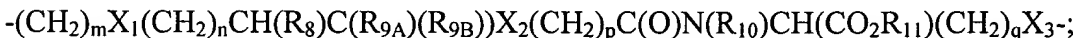
$\text{R}_{9\text{A}}$ and $\text{R}_{9\text{B}}$ together are oxo;

X_2 is NR_C ; and

B is hydrogen.

12 (Original). The compound according to claim 2, wherein

L is



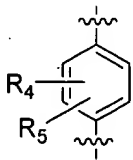
R_8 is $\text{NR}_\text{A}\text{R}_\text{B}$;

$\text{R}_{9\text{A}}$ and $\text{R}_{9\text{B}}$ together are oxo;

X_2 is NR_C ;

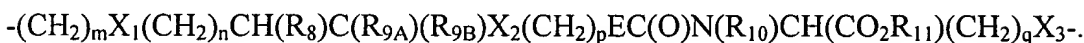
B is hydrogen; and

A is

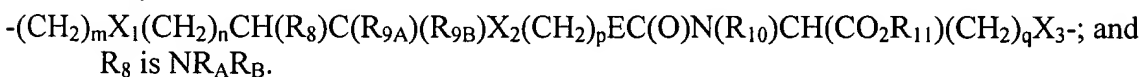


13 (Original). The compound according to claim 12, which is
N-[5-({N-acetyl-4-[(carboxycarbonyl)(2-carboxyphenyl)amino]-3-ethylphenylalanyl}amino)pentanoyl]-L-norleucine.

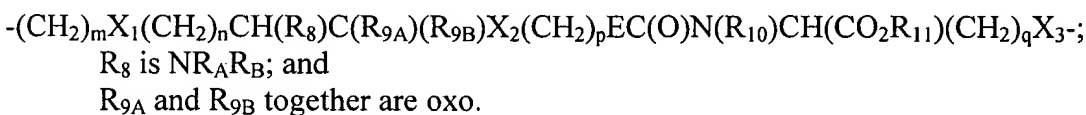
14 (Original). The compound according to claim 2, wherein
L is



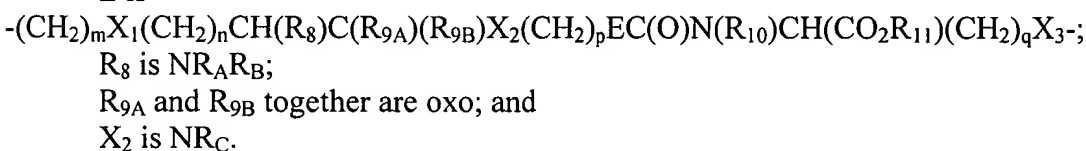
15 (Original). The compound according to claim 2, wherein
L is



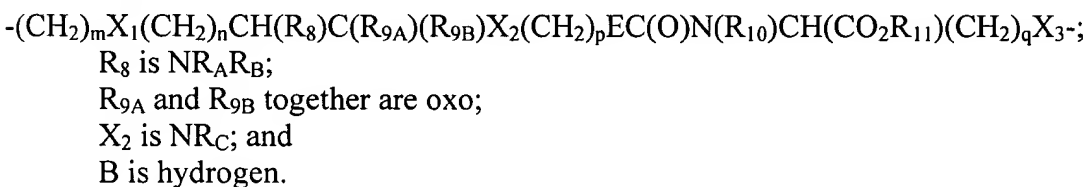
16 (Original). The compound according to claim 2, wherein
L is



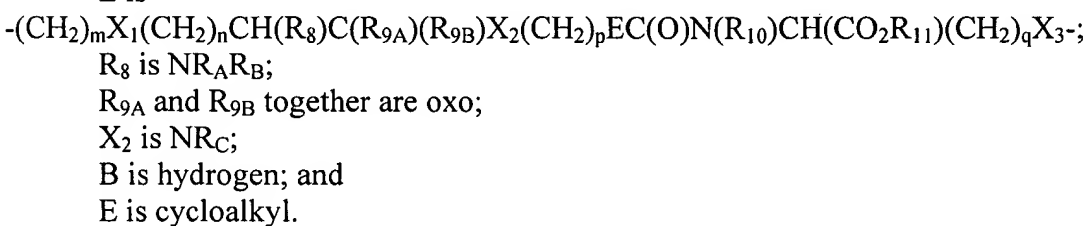
17 (Original). The compound according to claim 2, wherein
L is



18 (Original). The compound according to claim 2, wherein
L is



19 (Original). The compound according to claim 2, wherein
L is



20 (Original). The compound according to claim 2, wherein
L is

$-(CH_2)_mX_1(CH_2)_nCH(R_8)C(R_{9A})(R_{9B})X_2(CH_2)_pEC(O)N(R_{10})CH(CO_2R_{11})(CH_2)_qX_3-$;

R_8 is NR_AR_B ;

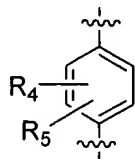
R_{9A} and R_{9B} together are oxo;

X_2 is NR_C ;

B is hydrogen;

E is cycloalkyl; and

A is



21 (Original). The compound according to claim 20, which is

$N-\{[4-\{[N\text{-acetyl-}4\text{-}[(\text{carboxycarbonyl})(2\text{-carboxyphenyl})\text{amino}]\text{-}3\text{-}(2\text{-hydroxyethyl})\text{phenylalanyl}]\text{amino}\}\text{methylcyclohexyl}]\text{carbonyl}\}\text{-L-norleucine}$.

22 (Original). The compound according to claim 2, wherein

L is

$-(CH_2)_mX_1(CH_2)_nCH(R_8)C(R_{9A})(R_{9B}))X_2(CH_2)_pC(O)N(R_{10})CH(CO_2R_{11})(CH_2)_qX_3-$;

R_8 is NR_AR_B ;

R_{9A} and R_{9B} together are oxo;

X_2 is NR_C ;

X_3 is S; and

B is alkyl.

23 (Original). The compound according to claim 2, wherein

L is

$-(CH_2)_mX_1(CH_2)_nCH(R_8)C(R_{9A})(R_{9B}))X_2(CH_2)_pC(O)N(R_{10})CH(CO_2R_{11})(CH_2)_qX_3-$;

R_8 is NR_AR_B ;

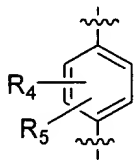
R_{9A} and R_{9B} together are oxo;

X_2 is NR_C ;

X_3 is S;

B is alkyl; and

A is



24 (Original). The compound according to claim 23, selected from the group consisting of

$N-\{5\text{-}[(N\text{-acetyl-}4\text{-}[(\text{carboxycarbonyl})(2\text{-carboxyphenyl})\text{amino}]\text{-}3\text{-ethylphenylalanyl})\text{amino}]\text{pentanoyl}\}\text{-L-methionine}$;

methyl $N-\{5\text{-}[(N\text{-acetyl-}4\text{-}[(\text{carboxycarbonyl})(2\text{-carboxyphenyl})\text{amino}]\text{-}3\text{-ethylphenylalanyl})\text{amino}]\text{pentanoyl}\}\text{-L-methioninate}$;

N-{5-[(*N*-acetyl-4-[(carboxycarbonyl)(2-carboxyphenyl)amino]-3-ethylphenylalanyl)amino]pentanoyl}-*S*-ethyl-L-homocysteine;

N-{5-[(*N*-acetyl-4-[(carboxycarbonyl)(2-carboxyphenyl)amino]-3-isopropylphenylalanyl)amino]pentanoyl}-L-methionine;

N-{5-[(*N*-acetyl-4-[(carboxycarbonyl)(2-carboxy-5-chlorophenyl)amino]-3-ethylphenylalanyl)amino]pentanoyl}-L-methionine; and

N-(5-{[*N*-acetyl-4-[(carboxycarbonyl)(2-carboxyphenyl)amino]-3-(2-hydroxyethyl)phenylalanyl]amino}pentanoyl)-L-methionine.

25 (Original). The compound according to claim 2, wherein

L is

$-(\text{CH}_2)_m\text{X}_1(\text{CH}_2)_n\text{CH}(\text{R}_8)\text{C}(\text{R}_{9A})(\text{R}_{9B})\text{X}_2(\text{CH}_2)_p\text{C}(\text{O})\text{N}(\text{R}_{10})\text{CH}(\text{CO}_2\text{R}_{11})(\text{CH}_2)_q\text{X}_3-$

R_8 is NR_AR_B ;

R_{9A} and R_{9B} together are oxo;

X_2 is NR_C ;

X_3 is S; and

B is aryl.

26 (Original). The compound according to claim 2, wherein

L is

$-(\text{CH}_2)_m\text{X}_1(\text{CH}_2)_n\text{CH}(\text{R}_8)\text{C}(\text{R}_{9A})(\text{R}_{9B})\text{X}_2(\text{CH}_2)_p\text{C}(\text{O})\text{N}(\text{R}_{10})\text{CH}(\text{CO}_2\text{R}_{11})(\text{CH}_2)_q\text{X}_3-$

R_8 is NR_AR_B ;

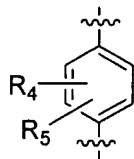
R_{9A} and R_{9B} together are oxo;

X_2 is NR_C ;

X_3 is S;

B is aryl; and

A is



27 (Original). The compound according to claim 26, which is

N-{5-[(*N*-acetyl-4-[(carboxycarbonyl)(2-carboxyphenyl)amino]-3-ethylphenylalanyl)amino]pentanoyl}-*S*-benzyl-L-cysteine.

28 (Original). The compound according to claim 2, wherein

L is

$-(\text{CH}_2)_m\text{X}_1(\text{CH}_2)_n\text{CH}(\text{R}_8)\text{C}(\text{R}_{9A})(\text{R}_{9B})\text{X}_2(\text{CH}_2)_p\text{C}(\text{O})\text{N}(\text{R}_{10})\text{CH}(\text{CO}_2\text{R}_{11})(\text{CH}_2)_q\text{X}_3-$

R_8 is NR_AR_B ;

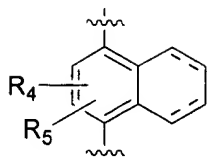
R_{9A} and R_{9B} together are oxo;

X_2 is NR_C ;

X_3 is S;

B is alkyl; and

A is



- 29 (Original). The compound according to claim 28, which is
N-(5-{[3-(4-[(carboxycarbonyl)(2-carboxyphenyl)amino]-1-naphthyl)-*N*-(methoxycarbonyl)alanyl]amino}pentanoyl)-*L*-methionine.
- 30 (Original). The compound according to claim 2, wherein
 L is $-(\text{CH}_2)_m\text{X}_1(\text{CH}_2)_n\text{CH}(\text{R}_8)\text{C}(\text{R}_{9A})(\text{R}_{9B})\text{X}_2(\text{CH}_2)_p\text{X}_3-$.
- 31 (Original). The compound according to claim 2, wherein
 L is $-(\text{CH}_2)_m\text{X}_1(\text{CH}_2)_n\text{CH}(\text{R}_8)\text{C}(\text{R}_{9A})(\text{R}_{9B})\text{X}_2(\text{CH}_2)_p\text{X}_3-$; and
 R_8 is NR_AR_B .
- 32 (Original). The compound according to claim 2, wherein
 L is $-(\text{CH}_2)_m\text{X}_1(\text{CH}_2)_n\text{CH}(\text{R}_8)\text{C}(\text{R}_{9A})(\text{R}_{9B})\text{X}_2(\text{CH}_2)_p\text{X}_3-$;
 R_8 is NR_AR_B ; and
 R_{9A} and R_{9B} together are oxo.
- 33 (Original). The compound according to claim 2, wherein
 L is $-(\text{CH}_2)_m\text{X}_1(\text{CH}_2)_n\text{CH}(\text{R}_8)\text{C}(\text{R}_{9A})(\text{R}_{9B})\text{X}_2(\text{CH}_2)_p\text{X}_3-$;
 R_8 is NR_AR_B ;
 R_{9A} and R_{9B} together are oxo; and
 X_2 is NR_C .
- 34 (Original). The compound according to claim 2, wherein
 L is $-(\text{CH}_2)_m\text{X}_1(\text{CH}_2)_n\text{CH}(\text{R}_8)\text{C}(\text{R}_{9A})(\text{R}_{9B})\text{X}_2(\text{CH}_2)_p\text{X}_3-$;
 R_8 is NR_AR_B ;
 R_{9A} and R_{9B} together are oxo;
 X_2 is NR_C ; and
 X_3 is O.
- 35 (Original). The compound according to claim 2, wherein
 L is $-(\text{CH}_2)_m\text{X}_1(\text{CH}_2)_n\text{CH}(\text{R}_8)\text{C}(\text{R}_{9A})(\text{R}_{9B})\text{X}_2(\text{CH}_2)_p\text{X}_3-$;
 R_8 is NR_AR_B ;
 R_{9A} and R_{9B} together are oxo;
 X_2 is NR_C ;
 X_3 is O; and
 B is aryl.

36 (Original). The compound according to claim 2, wherein

L is $-(\text{CH}_2)_m\text{X}_1(\text{CH}_2)_n\text{CH}(\text{R}_8)\text{C}(\text{R}_{9A})(\text{R}_{9B})\text{X}_2(\text{CH}_2)_p\text{X}_3-$;

R_8 is NR_AR_B ;

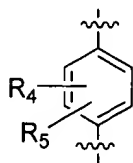
R_{9A} and R_{9B} together are oxo;

X_2 is NR_C ;

X_3 is O;

B is aryl; and

A is



37 (Original). The compound according to claim 36, selected from the group consisting of

- methyl 2-[4-({*N*-[(allyloxy)carbonyl]-4-[(carboxycarbonyl)(2-carboxyphenyl)amino]-*L*-phenylalanyl} amino)butoxy]-6-hydroxybenzoate;
- methyl 2-{4-[(*N*-acetyl-4-[(carboxycarbonyl)(2-carboxyphenyl)amino]-3-ethylphenylalanyl)amino]butoxy}-6-hydroxybenzoate;
- methyl 4-{4-[(*N*-acetyl-4-amino-3-ethylphenylalanyl)amino]butoxy}-2-hydroxy-1,1'-biphenyl-3-carboxylate;
- 2-[4-({*N*-acetyl-4-[(carboxycarbonyl)(2-carboxyphenyl)amino]-3-ethylphenylalanyl} amino)butoxy]-6-hydroxybenzoic acid;
- methyl 6-{4-[(*N*-acetyl-4-[(carboxycarbonyl)(2-carboxyphenyl)amino]-3-ethylphenylalanyl)amino]butoxy}-3-bromo-2-hydroxybenzoate;
- methyl 2-(4-{[4-[(carboxycarbonyl)(2-carboxyphenyl)amino]-*N*-(methoxycarbonyl)-*L*-phenylalanyl]amino}butoxy)-6-hydroxy-4-pentylbenzoate;
- methyl 2-(4-{[4-[(carboxycarbonyl)(2-carboxyphenyl)amino]-*N*-(methoxycarbonyl)-*L*-phenylalanyl]amino}butoxy)-6-hydroxy-4-methoxybenzoate;
- methyl 3-(4-{[4-[(carboxycarbonyl)(2-carboxyphenyl)amino]-*N*-(methoxycarbonyl)-*L*-phenylalanyl]amino}butoxy)-5-hydroxy-1,1'-biphenyl-4-carboxylate;
- methyl 2-(4-{[4-[(carboxycarbonyl)(2-carboxyphenyl)amino]-*N*-(methoxycarbonyl)-*L*-phenylalanyl]amino}butoxy)-6-hydroxy-4-methylbenzoate;
- methyl 2-(4-{[4-[(carboxycarbonyl)(2-carboxyphenyl)amino]-*N*-(methoxycarbonyl)-*L*-phenylalanyl]amino}butoxy)-4-chloro-6-hydroxybenzoate;
- methyl 2-(4-{[4-[(carboxycarbonyl)(2-carboxyphenyl)amino]-*N*-(methoxycarbonyl)-*L*-phenylalanyl]amino}butoxy)-6-hydroxybenzoate;
- 4-[(carboxycarbonyl)(2-carboxyphenyl)amino]-*N*-{4-[2-(aminocarbonyl)-3-hydroxyphenoxy]butyl}-*N*-(methoxycarbonyl)-*L*-phenylalaninamide;
- methyl 3-(4-{[4-[(carboxycarbonyl)(2-carboxyphenyl)amino]-*N*-(methoxycarbonyl)-*L*-phenylalanyl]amino}butoxy)-1-hydroxy-2-naphthoate;
- 4-[(carboxycarbonyl)(2-carboxyphenyl)amino]-*N*-(4-{3-hydroxy-2-[(methylamino)carbonyl]phenoxy}butyl)-*N*-(methoxycarbonyl)-*L*-phenylalaninamide;

4-[(carboxycarbonyl)(2-carboxyphenyl)amino]-*N*-(4-{2-[(ethylamino)carbonyl]-3-hydroxyphenoxy}butyl)-*N*-(methoxycarbonyl)-L-phenylalaninamide;
N-{4-[2-(acetylamino)-3-hydroxyphenoxy]butyl}-4-[(carboxycarbonyl)(2-carboxyphenyl)amino]-*N*-(methoxycarbonyl)-L-phenylalaninamide; and
 4-[(carboxycarbonyl)(2-carboxyphenyl)amino]-*N*-(4-{2-[(dimethylamino)carbonyl]-3-hydroxyphenoxy}butyl)-*N*-(methoxycarbonyl)-L-phenylalaninamide.

38 (Original). The compound according to claim 2, wherein

L is $-(\text{CH}_2)_m\text{X}_1(\text{CH}_2)_n\text{CH}(\text{R}_8)\text{C}(\text{R}_{9A})(\text{R}_{9B})\text{X}_2(\text{CH}_2)_p\text{X}_3-$;

R₈ is NR_AR_B;

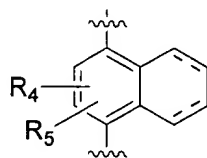
R_{9A} and R_{9B} together are oxo;

X₂ is NR_C;

X₃ is O;

B is aryl; and

A is



39 (Original). The compound according to claim 38, selected from the group consisting of

methyl 2-[(5-{[*N*-acetyl-3-(4-amino-1-naphthyl)-L-alanyl]amino}pentyl)oxy]-6-hydroxy-4-methylbenzoate; and

3-({5-[*N*-acetyl-3-{4-[(carboxycarbonyl)(2-carboxyphenyl)amino]-1-naphthyl}-L-alanyl]amino}pentyl)oxy)-2-naphthoic acid.

40 (Original). The compound according to claim 2, wherein

L is $-(\text{CH}_2)_m\text{X}_1(\text{CH}_2)_n\text{CH}(\text{R}_8)\text{C}(\text{R}_{9A})(\text{R}_{9B})\text{X}_2(\text{CH}_2)_p\text{X}_3-$; and

R₈ is hydrogen.

41 (Original). The compound according to claim 2, wherein

L is $-(\text{CH}_2)_m\text{X}_1(\text{CH}_2)_n\text{CH}(\text{R}_8)\text{C}(\text{R}_{9A})(\text{R}_{9B})\text{X}_2(\text{CH}_2)_p\text{X}_3-$;

R₈ is hydrogen; and

R_{9A} and R_{9B} together are oxo.

42 (Original). The compound according to claim 2, wherein

L is $-(\text{CH}_2)_m\text{X}_1(\text{CH}_2)_n\text{CH}(\text{R}_8)\text{C}(\text{R}_{9A})(\text{R}_{9B})\text{X}_2(\text{CH}_2)_p\text{X}_3-$;

R₈ is hydrogen;

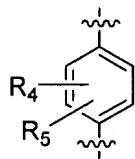
R_{9A} and R_{9B} together are oxo; and

X₂ is NR_C.

43 (Original). The compound according to claim 2, wherein
 L is $-(\text{CH}_2)_m\text{X}_1(\text{CH}_2)_n\text{CH}(\text{R}_8)\text{C}(\text{R}_{9A})(\text{R}_{9B})\text{X}_2(\text{CH}_2)_p\text{X}_3-$;
 R_8 is hydrogen;
 R_{9A} and R_{9B} together are oxo;
 X_2 is NR_C ; and
 X_3 is O.

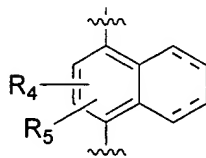
44 (Original). The compound according to claim 2, wherein
 L is $-(\text{CH}_2)_m\text{X}_1(\text{CH}_2)_n\text{CH}(\text{R}_8)\text{C}(\text{R}_{9A})(\text{R}_{9B})\text{X}_2(\text{CH}_2)_p\text{X}_3-$;
 R_8 is hydrogen;
 R_{9A} and R_{9B} together are oxo;
 X_2 is NR_C ;
 X_3 is O; and
 B is aryl.

45 (Original). The compound according to claim 2, wherein
 L is $-(\text{CH}_2)_m\text{X}_1(\text{CH}_2)_n\text{CH}(\text{R}_8)\text{C}(\text{R}_{9A})(\text{R}_{9B})\text{X}_2(\text{CH}_2)_p\text{X}_3-$;
 R_8 is hydrogen;
 R_{9A} and R_{9B} together are oxo;
 X_2 is NR_C ;
 X_3 is O; and
 B is aryl; and
 A is



46 (Original). The compound according to claim 45, which is
 methyl 2-(4-{[3-(4-[(carboxycarbonyl)(2-carboxyphenyl)amino]-3-ethylphenyl)propanoyl]amino}butoxy)-6-hydroxybenzoate.

47 (Original). The compound according to claim 2, wherein
 L is $-(\text{CH}_2)_m\text{X}_1(\text{CH}_2)_n\text{CH}(\text{R}_8)\text{C}(\text{R}_{9A})(\text{R}_{9B})\text{X}_2(\text{CH}_2)_p\text{X}_3-$;
 R_8 is hydrogen;
 R_{9A} and R_{9B} together are oxo;
 X_2 is NR_C ;
 X_3 is O;
 B is aryl; and
 A is



48 (Original). The compound according to claim 47, which is

2-((carboxycarbonyl){4-[3-({4-[3-hydroxy-2-(methoxycarbonyl)phenoxy]butyl}amino)-3-oxopropyl]-[(carboxycarbonyl)(2-carboxyphenyl)amino]-1-naphthyl}amino)benzoic acid.

49 (Original). The compound according to claim 2, wherein

L is $-(\text{CH}_2)_m\text{X}_1(\text{CH}_2)_n\text{CH}(\text{R}_8)\text{C}(\text{R}_{9A})(\text{R}_{9B})\text{X}_2(\text{CH}_2)_p\text{X}_3$;

R_8 is hydrogen; and

R_{9A} is alkyl.

50 (Original). The compound according to claim 2, wherein

L is $-(\text{CH}_2)_m\text{X}_1(\text{CH}_2)_n\text{CH}(\text{R}_8)\text{C}(\text{R}_{9A})(\text{R}_{9B})\text{X}_2(\text{CH}_2)_p\text{X}_3$;

R_8 is hydrogen;

R_{9A} is alkyl; and

X_2 is NR_C .

51 (Original). The compound according to claim 2, wherein

L is $-(\text{CH}_2)_m\text{X}_1(\text{CH}_2)_n\text{CH}(\text{R}_8)\text{C}(\text{R}_{9A})(\text{R}_{9B})\text{X}_2(\text{CH}_2)_p\text{X}_3$;

R_8 is hydrogen;

R_{9A} is alkyl;

X_2 is NR_C ; and

X_3 is O.

52 (Original). The compound according to claim 2, wherein

L is $-(\text{CH}_2)_m\text{X}_1(\text{CH}_2)_n\text{CH}(\text{R}_8)\text{C}(\text{R}_{9A})(\text{R}_{9B})\text{X}_2(\text{CH}_2)_p\text{X}_3$;

R_8 is hydrogen;

R_{9A} is alkyl;

X_2 is NR_C ;

X_3 is O; and

B is aryl.

53 (Original). The compound according to claim 2, wherein

L is $-(\text{CH}_2)_m\text{X}_1(\text{CH}_2)_n\text{CH}(\text{R}_8)\text{C}(\text{R}_{9A})(\text{R}_{9B})\text{X}_2(\text{CH}_2)_p\text{X}_3$;

R_8 is hydrogen;

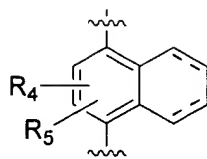
R_{9A} is alkyl;

X_2 is NR_C ;

X_3 is O;

B is aryl; and

A is



54 (Original). The compound according to claim 53, which is methyl 2-(4-{{3-(4-[(carboxycarbonyl)(2-carboxyphenyl)amino]-1-naphthyl)-1-methylpropyl}amino}butoxy)-6-hydroxybenzoate.

55 (Original). The compound according to claim 2, wherein
 L is $-(\text{CH}_2)_m\text{X}_1(\text{CH}_2)_n\text{CH}(\text{R}_8)\text{C}(\text{R}_{9A})(\text{R}_{9B})\text{X}_2(\text{CH}_2)_p\text{X}_3-$;
 R_8 is hydrogen; and
 R_{9A} and R_{9B} are both hydrogen.

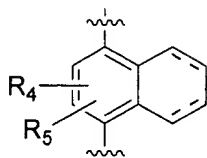
56 (Original). The compound according to claim 2, wherein
 L is $-(\text{CH}_2)_m\text{X}_1(\text{CH}_2)_n\text{CH}(\text{R}_8)\text{C}(\text{R}_{9A})(\text{R}_{9B})\text{X}_2(\text{CH}_2)_p\text{X}_3-$;
 R_8 is hydrogen;
 R_{9A} and R_{9B} are both hydrogen; and
 X_2 is NR_C .

57 (Original). The compound according to claim 2, wherein
 L is $-(\text{CH}_2)_m\text{X}_1(\text{CH}_2)_n\text{CH}(\text{R}_8)\text{C}(\text{R}_{9A})(\text{R}_{9B})\text{X}_2(\text{CH}_2)_p\text{X}_3-$;
 R_8 is hydrogen;
 R_{9A} and R_{9B} are both hydrogen;
 X_2 is NR_C ; and
 X_3 is O.

58 (Original). The compound according to claim 2, wherein
 L is $-(\text{CH}_2)_m\text{X}_1(\text{CH}_2)_n\text{CH}(\text{R}_8)\text{C}(\text{R}_{9A})(\text{R}_{9B})\text{X}_2(\text{CH}_2)_p\text{X}_3-$;
 R_8 is hydrogen;
 R_{9A} and R_{9B} are both hydrogen;
 X_2 is NR_C ;
 X_3 is O; and
 B is aryl.

59 (Original). The compound according to claim 2, wherein
 L is $-(\text{CH}_2)_m\text{X}_1(\text{CH}_2)_n\text{CH}(\text{R}_8)\text{C}(\text{R}_{9A})(\text{R}_{9B})\text{X}_2(\text{CH}_2)_p\text{X}_3-$;
 R_8 is hydrogen;
 R_{9A} and R_{9B} are both hydrogen;
 X_2 is NR_C ;

X_3 is O;
 B is aryl; and
 A is



60 (Original). The compound according to claim 59, which is methyl 2-(4-{[3-(4-[(carboxycarbonyl)(2-carboxyphenyl)amino]-1-naphthyl)propyl]amino}butoxy)-6-hydroxybenzoate.

61 (Original). The compound according to claim 2, wherein

L is $-(CH_2)_mX_1(CH_2)_nCH(R_8)C(R_{9A})(R_{9B})X_2(CH_2)_pX_3(CH_2)_qX_4-$.

62 (Original). The compound according to claim 2, wherein

L is $-(CH_2)_mX_1(CH_2)_nCH(R_8)C(R_{9A})(R_{9B})X_2(CH_2)_pX_3(CH_2)_qX_4-$; and
 R_8 is NR_AR_B .

63 (Original). The compound according to claim 2, wherein

L is $-(CH_2)_mX_1(CH_2)_nCH(R_8)C(R_{9A})(R_{9B})X_2(CH_2)_pX_3(CH_2)_qX_4-$;
 R_8 is NR_AR_B ; and
 R_{9A} and R_{9B} together are oxo.

64 (Original). The compound according to claim 2, wherein

L is $-(CH_2)_mX_1(CH_2)_nCH(R_8)C(R_{9A})(R_{9B})X_2(CH_2)_pX_3(CH_2)_qX_4-$;
 R_8 is NR_AR_B ;
 R_{9A} and R_{9B} together are oxo; and
 X_2 is NR_C .

65 (Original). The compound according to claim 2, wherein

L is $-(CH_2)_mX_1(CH_2)_nCH(R_8)C(R_{9A})(R_{9B})X_2(CH_2)_pX_3(CH_2)_qX_4-$;
 R_8 is NR_AR_B ;
 R_{9A} and R_{9B} together are oxo;
 X_2 is NR_C ; and
 X_3 is O.

66 (Original). The compound according to claim 2, wherein

L is $-(CH_2)_mX_1(CH_2)_nCH(R_8)C(R_{9A})(R_{9B})X_2(CH_2)_pX_3(CH_2)_qX_4-$;
 R_8 is NR_AR_B ;

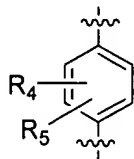
R_{9A} and R_{9B} together are oxo;
 X_2 is NR_C ;
 X_3 is O; and
 X_4 is O.

67 (Original). The compound according to claim 2, wherein

L is $-(CH_2)_mX_1(CH_2)_nCH(R_8)C(R_{9A})(R_{9B})X_2(CH_2)_pX_3(CH_2)_qX_4-$;
 R_8 is NR_AR_B ;
 R_{9A} and R_{9B} together are oxo;
 X_2 is NR_C ;
 X_3 is O;
 X_4 is O; and
 B is aryl.

68 (Original). The compound according to claim 2, wherein

L is $-(CH_2)_mX_1(CH_2)_nCH(R_8)C(R_{9A})(R_{9B})X_2(CH_2)_pX_3(CH_2)_qX_4-$;
 R_8 is NR_AR_B ;
 R_{9A} and R_{9B} together are oxo;
 X_2 is NR_C ;
 X_3 is O;
 X_4 is O;
 B is aryl; and
 A is



69 (Original). The compound according to claim 68, which is

methyl 2-{2-[2-({N-[(allyloxy)carbonyl]-4-[(carboxycarbonyl)(2-carboxyphenyl)amino]-L-phenylalanyl}amino)ethoxy]ethoxy}-6-hydroxybenzoate;

70 (Original). A pharmaceutical composition comprising a therapeutically effective amount of a compound of claim 1 in combination with a pharmaceutically acceptable carrier.

71 (Currently Amended). A method ~~of method~~ of treating diabetes by selectively inhibiting protein tyrosine phosphatase 1B comprising administering a therapeutically effective amount of a compound of claim 1.

72 (Currently Amended). A method of treating diabetes disorders caused by overexpressed or altered protein tyrosine phosphatase 1B comprising administering a therapeutically effective amount of a compound of claim 1.

73 (Canceled). The method of claim 72, wherein the disorder is type I and type II diabetes.

74 (Canceled). The method of claim 72, wherein the disorder is obesity.

75 (Canceled). A method of claim 72, wherein the disorder is autoimmune disorders, acute and chronic inflammatory disorders, osteoporosis, cancer, malignant disorders.

RESPONSE

Applicants have amended the claims to reflect the election of group I made on 2/21/03. All claimed limitations related to heterocyclic compounds have been removed from the claims. Applicants respectfully maintain that claims 1-70 are now in condition for allowance.

Claims 71-75 stand rejected under 35 U.S.C. 112, first paragraph. Examiner maintains that the specification is not enabling for the claims directed to treating disorders caused by overexpressed or altered protein tyrosine phosphatase 1B. The Examiner maintains that the claims are not directed to specific diseases, but all kinds of diseases that use the mechanistic nature of inhibiting protein tyrosine phosphatase 1B. Applicants respectfully traverse this rejection and request withdrawal of the same.

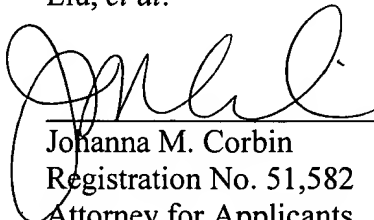
Applicants have amended claims 71 and 72 to specify a disease treatable by inhibiting protein tyrosine phosphatase 1B. Claims 73-75 have been canceled. Applicants respectfully maintain that amended claims 71 and 72 are now in condition for allowance.

ACTION REQUESTED

For all the forgoing reasons, Applicants submit that Claims 1-72 are in condition for allowance. To that end, the examiner is invited to contact the undersigned to schedule an Examiner Interview to discuss any matter.

Respectfully submitted,
Liu, *et al.*

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